



4

SEQUENCE LISTING

<110> Bowdish, Katherine S.
Frederickson, Shana
Renshaw, Mark

<120> RATIONALLY DESIGNED ANTIBODIES

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<140> 10/006,593
<141> 2001-12-05

<150> US 60/251,448
<151> 2000-12-05

<150> US 60/288,889
<151> 2001-05-04

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<151> 2001-05-29

<160> 118

<170> PatentIn version 3.1

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Gly Gly

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<210> 9
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<220>
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<210> 19
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<210> 25
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<400> 25

Pro Pro Ile Glu Gly Pro Thr Leu Arg Gln Trp Leu Ala Ala Arg Ala
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Gly Gly

<210> 26
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<400> 27

Gly Gly Ile Glu Gly Pro Thr Leu Arg Gln Trp Leu Ala Ala Arg Ala
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Gly Gly

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Gly Gly

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Pro Val

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Val Gly

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Pro Asp

<210> 38
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54

<210> 39
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<220>
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<400> 39

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1 5 10 15

Pro Val

<210> 40
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<220>
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<210> 41
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<220>
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<400> 41

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1 5 10 15

Pro Ile

<210> 42
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1 5 10 15

Pro Val

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1 5 10 15

Pro Val

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<212> DNA
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Cys Ser

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1 5 10 15

Pro Asp

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ctgcgcgaac aggtggcaca gctgaaacag aaagttatga accatggcg ggttgtctag      180
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Arg Leu Glu Glu Lys Val Lys Thr Leu Lys Ala Gln Asn Ser Glu Leu
20 25 30

Ala Ser Thr Ala Asn Met Leu Arg Glu Gln Val Ala Gln Leu Lys Gln
35 40 45

Lys Val Met Asn His Gly Gly Cys Ala Ser Gly Gln Ala Gly Gln His
50 55 60

His His His His Gly Ala Tyr Pro Tyr Asp Val Pro Asp Tyr Ala
65 70 75 80

Ser

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gaggccaaAG tacagtggAA ggtggataAC gccctccaAT cgggtAACTC ccaggagAGT      480
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aaAGCAGACT acgagAAAACA caaaGTCTAC gcCTGCGAAG tcACCCATCA gggcCTgAGC      600
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1 5 10 15

Leu Ala Ala Arg Ala Xaa Xaa Trp Gly Gln Gly Thr
20 25

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<220>
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agatagggtt gagtgttgc ccagttgga acaagagtcc actattaaag aacgtggact      180
ccaacgtcaa agggcgaaaa accgtctatc agggcgatgg cccactacgt gaaccatcac      240

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aagcgaaagg	agcgggcgct	agggcgctgg	caagtgtac	ggtcacgctg	cgcgttaacca	420
ccacacccgc	cgcgttaat	gcgcgcgtac	agggcgcgtc	aggtggcact	tttcggggaa	480
atgtgcgcgg	aacccttatt	tgtttatttt	tctaaataca	ttcaaataatg	tatccgtca	540
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<210> 61
 <211> 18
 <212> PRT
 <213> artificial sequence.

<220>

<223> TPO mimetic with flanking amino acids

<400> 61

Asn Pro Ile Glu Gly Pro Thr Leu Arg Gln Trp Leu Ala Ala Arg Ala
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Arg Gly

<210> 62

<211> 41

<212> DNA

<213> artificial sequence

<220>

<223> primer

<400> 62

taggatgcgg ccgcacaggt cttttttttt tttttttttt t

41

<210> 63

<211> 24

<212> DNA

<213> artificial sequence

<220>

<223> primer

<400> 63

ccatgttaggc tgtgcccgtg gatt

24

<210> 64

<211> 24

<212> DNA

<213> artificial sequence

<220>

<223> primer

<400> 64

ccacgggcac agcctacatg gagc

24

<210> 65

<211> 54

<212> DNA

<213> artificial sequence

<220>

<223> nucleic acid encoding TPO mimetic peptide flanking sequence

<400> 65

ttgccaattg aagggccgac gctgcggcaa tggctggcgg cgcgccgcgc ttgtt

54

<210> 66
<211> 18
<212> PRT
<213> artificial sequence

<220>
<223> TPO mimetic peptide with flanking sequence

<400> 66

Leu Pro Ile Glu Gly Pro Thr Leu Arg Gln Trp Leu Ala Ala Arg Ala
1 5 10 15

Pro Val

<210> 67
<211> 472
<212> PRT
<213> artificial sequence

<220>
<223> Humanized antibody heavy chain

<400> 67

Met Lys Trp Ser Trp Val Ile Leu Phe Leu Leu Ser Val Thr Ala Gly
1 5 10 15

Val His Ser Gln Val Gln Leu Val Gln Ser Gly Ala Glu Val Lys Lys
20 25 30

Pro Gly Ala Ser Val Lys Val Ser Cys Lys Ala Ser Gly Tyr Ile Phe
35 40 45

Ser Asn Tyr Trp Ile Gln Trp Val Arg Gln Ala Pro Gly Gln Gly Leu
50 55 60

Glu Trp Met Gly Glu Ile Leu Pro Gly Ser Gly Ser Thr Glu Tyr Thr
65 70 75 80

Glu Asn Phe Lys Asp Arg Val Thr Met Thr Arg Asp Thr Ser Thr Ser
85 90 95

Thr Val Tyr Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Val
100 105 110

Tyr Tyr Cys Ala Arg Leu Pro Ile Glu Gly Pro Thr Leu Arg Gln Trp
115 120 125

Leu Ala Ala Arg Ala Pro Val Trp Gly Gln Gly Thr Leu Val Thr Val
130 135 140

Ser Ser Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Cys
145 150 155 160

Ser Arg Ser Thr Ser Glu Ser Thr Ala Ala Leu Gly Cys Leu Val Lys
165 170 175

Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu
180 185 190

Thr Ser Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu
195 200 205

Tyr Ser Leu Ser Ser Val Val Thr Val Pro Ser Ser Asn Phe Gly Thr
210 215 220

Gln Thr Tyr Thr Cys Asn Val Asp His Lys Pro Ser Asn Thr Lys Val
225 230 235 240

Asp Lys Thr Val Glu Arg Lys Cys Cys Val Glu Cys Pro Pro Cys Pro
245 250 255

Ala Pro Pro Val Ala Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro
260 265 270

Lys Asp Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val
275 280 285

Val Asp Val Ser Gln Glu Asp Pro Glu Val Gln Phe Asn Trp Tyr Val
290 295 300

Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln
305 310 315 320

Phe Asn Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln
325 330 335

Asp Trp Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Gly
340 345 350

Leu Pro Ser Ser Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro
355 360 365

Arg Glu Pro Gln Val Tyr Thr Leu Pro Pro Ser Gln Glu Glu Met Thr
370 375 380

Lys Asn Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser
385 390 395 400

Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr
405 410 415

Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr
420 425 430

Ser Arg Leu Thr Val Asp Lys Ser Arg Trp Gln Glu Gly Asn Val Phe
435 440 445

Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys
450 455 460

Ser Leu Ser Leu Ser Leu Gly Lys
465 470

<210> 68
<211> 1419
<212> DNA
<213> artificial sequence

<220>
<223> nucleic acid encoding humanized antibody heavy chain

<400> 68
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tgtaaagcta gcggctataat tttttctaat tattggattc aatgggtgcg tcaggcccc 180
gggcagggcc tggaaatggat gggtgagatc ttaccgggct ctggtagcac cgaatatacc 240
gaaaatttta aagaccgtgt tactatgacg cgtgacactt cgactagttac agtatacatg 300
gagctctcca gcctgcgatc ggaggacacg gccgtctatt attgcgcg 360

gaaggggccga	cgctgcggca	atggctggcg	gchgccccgc	ctgtttgggg	tcaaggaacc	420
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aggtggcagg	agggaaatgt	cttctcatgc	tccgtgatgc	atgaggctct	gcacaaccac	1380
tacacacaga	agagcctctc	cctgtctctg	ggtaaatga			1419

<210> 69
 <211> 236
 <212> PRT
 <213> artificial sequence

<220>
 <223> Humanized antibody light chain

<400> 69

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20															30

Leu Ser Ala Ser Val Gly Asp Arg Val Thr Ile Thr Cys Gly Ala Ser
35 40 45

Glu Asn Ile Tyr Gly Ala Leu Asn Trp Tyr Gln Gln Lys Pro Gly Lys
50 55 60

Ala Pro Lys Leu Leu Ile Tyr Gly Ala Thr Asn Leu Ala Asp Gly Val
65 70 75 80

Pro Ser Arg Phe Ser Gly Ser Gly Thr Asp Phe Thr Leu Thr
85 90 95

Ile Ser Ser Leu Gln Pro Glu Asp Phe Ala Thr Tyr Tyr Cys Gln Asn
100 105 110

Val Leu Asn Thr Pro Leu Thr Phe Gly Gln Gly Thr Lys Val Glu Ile
115 120 125

Lys Arg Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp
130 135 140

Glu Gln Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn
145 150 155 160

Phe Tyr Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu
165 170 175

Gln Ser Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp
180 185 190

Ser Thr Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr
195 200 205

Glu Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser
210 215 220

Ser Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
225 230 235

<210> 70
<211> 711
<212> DNA
<213> artificial sequence

<220>
<223> nucleic acid encoding humanized antibody light chain

<400> 70
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gtcaccatca cctgcggcgc cagcgaaaac atctatggcg cgctgaactg gtatcaacag 180
aaacccggaa aagctccgaa gcttctgatt tacggtgcga cgaacctggc agatggagtc 240
ccttctcgct tctctggatc cggtccggaa acggattca ctctgaccat cagcagtctg 300
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ccgcacatctg atgagcagtt gaaatctgga actgcctctg ttgtgtgcct gctgaataac 480
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tcccaggaga gtgtcacaga gcaggacagc aaggacagca cctacagcct cagcagcacc 600
ctgacgctga gcaaagcaga ctacgagaaa cacaaagtct acgcctgcga agtcacccat 660
caggcctga gtcgcccgt cacaaagagc ttcaacaggg gagagtgtta g 711

<210> 71
<211> 22
<212> PRT
<213> artificial sequence

<220>
<223> EPO mimetic with random flanking amino acids

<220>
<221> MISC_FEATURE
<222> (1)..(2)
<223> Xaa is any amino acid

<220>
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<222> (6)..(6)
<223> Xaa is any amino acid

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<223> Xaa is any amino acid

<220>

<221> MISC_FEATURE
<222> (21)..(22)
<223> Xaa is any amino acid

<400> 71

Xaa Xaa Asp Tyr His Xaa Arg Met Gly Pro Leu Thr Trp Val Xaa Lys
1 5 10 15

Pro Leu Gly Gly Xaa Xaa
20

<210> 72
<211> 21
<212> DNA
<213> artificial sequence

<220>
<223> primer

<400> 72
taggatgcgg ccgcacaggt c

21

<210> 73
<211> 39
<212> DNA
<213> artificial sequence

<220>
<223> primer

<400> 73
cacgcgcaca acacgtctag aracatccag atgaccagg

39

<210> 74
<211> 39
<212> DNA
<213> artificial sequence

<220>
<223> primer

<400> 74
cacgcgcaca acacgtctag agmcatccag ttgaccagg

39

<210> 75
<211> 39
<212> DNA
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<220>

<223> primer

<400> 75
cacgcgcaca acacgtctag agccatccrg atgaccagg 39

<210> 76
<211> 39
<212> DNA
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<220>
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<400> 76
cacgcgcaca acacgtctag agtcatctgg atgaccagg 39

<210> 77
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<220>
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<400> 77
cacgcgcaca acacgtctag agatattgtg atgaccagg 39

<210> 78
<211> 39
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<220>
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<400> 78
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<210> 79
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<220>
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<400> 79
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<210> 80
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<400> 81
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<210> 82
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<400> 82
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<210> 83
<211> 39
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<220>
<223> primer

<400> 83
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<210> 84
<211> 39
<212> DNA
<213> artificial sequence

<220>
<223> primer

<400> 84
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<223> primer		
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<220>		
<223> primer		
<400> 86		
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<211> 20		
<212> DNA		
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<223> primer		
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<211> 59		
<212> DNA		
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<220>
 <223> part of mimetic

<400> 112

Gly Pro Thr Leu Arg Gln Trp Leu
1 5

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<222> (2)..(2)
<223> Xaa is any amino acid
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<220>
<221> MISC_FEATURE
<222> (11)..(11)
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<400> 113
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Gln Gly
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<223> primer
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```
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<221> misc_feature
<222> (26)..(27)
<223> n is a, c, g or t
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```
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<220>
<223> primer
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```
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<221> misc_feature
<222> (25)..(26)
<223> n is a, c, g or t
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ggaaagctc ctaagctcct gatctataac ccgatcgaag gcccaaccct gcgccagtg 180
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gatttcactc tcaccatcag cagcctgcag cctgaagatt ttgcaactta ttactgccaa 300
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<213> human

<400> 117

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Ser Ser Val Lys Val Ser Cys Arg Ala Ser Gly Gly Thr Phe Asn Asn
20 25 30

Tyr Ala Ile Ser Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp
35 40 45

Met Gly Gly Ile Phe Pro Phe Arg Asn Thr Ala Lys Tyr Ala Gln His
50 55 60

Phe Gln Gly Arg Val Thr Ile Thr Ala Asp Glu Ser Thr Gly Thr Ala
65 70 75 80

Tyr Met Glu Leu Ser Ser Leu Arg Ser Glu Asp Thr Ala Ile Tyr Tyr
85 90 95

Cys Ala Arg Gly Asp Thr Ile Phe Gly Val Thr Met Gly Tyr Tyr Ala
100 105 110

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Met Asp Val Trp Gly Gln Gly Thr Thr Val Thr Val Ser Ala Ala Ser
115 120 125

Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Ser Ser Lys Ser Thr
130 135 140

Ser Gly Gly Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro
145 150 155 160

Glu Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu Thr Ser Gly Val
165 170 175

His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser Leu Ser
180 185 190

Ser Val Val Thr Val Pro Ser Ser Ser Leu Gly Thr Gln Thr Tyr Ile
195 200 205

Cys Asn Val Asn His Lys Pro Ser Asn Thr Lys Val Asp Lys Lys Val
210 215 220

Glu Pro Lys Ser Cys Asp Lys Thr Ser
225 230

<210> 118
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<213> human

<400> 118

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Ala Thr Leu Ser Cys Arg Ala Ser His Ser Val Ser Arg Ala Tyr Leu
20 25 30

Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro Arg Leu Leu Ile Tyr
35 40 45

Gly Thr Ser Ser Arg Ala Thr Gly Ile Pro Asp Arg Phe Ser Gly Ser
50 55 60

Gly Ser Gly Thr Asp Phe Thr Leu Thr Ile Ser Arg Leu Glu Pro Glu
65 70 75 80

Asp Phe Ala Val Tyr Tyr Cys Gln Gln Tyr Gly Gly Ser Pro Trp Phe
85 90 95

Gly Gln Gly Thr Lys Val Glu Leu Lys Arg Thr Val Ala Ala Pro Ser
100 105 110

Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys Ser Gly Thr Ala
115 120 125

Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg Glu Ala Lys Val
130 135 140

Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly Asn Ser Gln Glu Ser
145 150 155 160

Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr Ser Leu Ser Ser Thr
165 170 175

Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His Lys Val Tyr Ala Cys
180 185 190

Glu Val Thr His Gln Gly Leu Ser Leu Pro Val Thr Lys Ser Phe Asn
195 200 205

Arg Gly Glu Cys
210